AMENDMENTS TO THE CLAIMS

The following listing of claims will replace all prior versions, and listings, of claims in the application.

Listing of claims:

1 and 2 (canceled).

3 (currently amended). The method of claim 2 A method for treating chronic pain, wherein said chronic pain is a type of neuropathic pain, said method comprising administering to a subject in need of such treatment a composition comprising a MEK inhibitor selected from a compound of the following formula (I):

$$\mathbf{W} \stackrel{\mathbf{O}}{\longrightarrow} \mathbf{R}_{11} \stackrel{\mathbf{H}}{\longrightarrow} \mathbf{R}_{10}$$

wherein

W is OR_1 , NR_2OR_1 , NR_AR_B , $NR_2NR_AR_B$, $O(CH_2)_{2-4}NR_AR_B$, or $NR_2(CH_2)_{2-4}$ NR_AR_B ;

R₁ is H, C₁₋₈ alkyl, C₃₋₈ alkenyl, C₃₋₈ alkynyl, C₃₋₈ cycloalkyl, phenyl, (phenyl)-C₁₋₄ alkyl, (phenyl)C₃₋₄ alkenyl, (phenyl)C₃₋₄ alkynyl, (C₃₋₈ cycloalkyl)C₁₋₄ alkyl, (C₃₋₈ cycloalkyl)C₃₋₄ alkenyl, (C₃₋₈ cycloalkyl)C₃₋₄ alkynyl, C₃₋₈ heterocyclic radical)C₁₋₄ alkyl, (C₃₋₈ heterocyclic radical)C₃₋₄ alkynyl or (CH₂)₂₋₄ NR_CR_D;

R₂ is H, C₁₋₄ alkyl, phenyl, C₃₋₆ cycloalkyl, C₃₋₆ heterocyclic radical, or (C₃₋₆ cycloalkyl) methyl;

R_A is H, C ₁₋₆ alkyl, C ₃₋₈ alkenyl, C ₃₋₈ alkynyl, C ₃₋₈ cycloalkyl, phenyl, (C ₃₋₈ cycloalkyl)C ₁₋₄ alkyl, (C ₃₋₈ cycloalkyl)C ₃₋₄ alkenyl, (C ₃₋₈ cycloalkyl)C ₃₋₄ alkynyl, C ₃₋₈ heterocyclic radical, (C ₃₋₈ heterocyclic radical)C ₁₋₄ alkyl, (aminosulfonyl)phenyl, [(aminosulfonyl)phenyl]C ₁₋₄ alkyl, (aminosulfonyl)C ₁₋₆ alkyl, (aminosulfonyl)C ₃₋₆ cycloalkyl, [(aminosulfonyl)C ₃₋₆ cycloalkyl]C ₁₋₄ alkyl, or (CH₂)₂₋₄ NR_CR_D;

R_B is H, C₁₋₈ alkyl, C₃₋₈ alkenyl, C₃₋₈ alkynyl, C₃₋₈ cycloalkyl, or phenyl;

Q is one of the following formulae (i) – (iii):

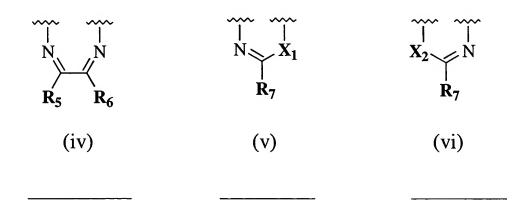
$$R_3$$
 R_4
 R_4

 R_3 is H or F;

R4 is halo, NO₂, SO₂NR_O(CH₂)_{2.4}NR_ER_F, SO₂NR_ER_F, or (CO)T;

T is C₁₋₈ alkyl, C₃₋₈ cycloalkyl, (NR_ER_F)C₁₋₄ alkyl, OR_F, -NR_O(CH₂)₂₋₄ NR_ER_F, or NR_ER_F;

Z is one of the following formulae (iv) – (viii):



one of R_5 and R_6 is H or methyl and the other of R_5 and R_6 is H, C_{1-6} alkyl, C_{2-6} alkenyl, C_{2-6} alkynyl, phenyl, benzyl, or -M-E-G;

 \underline{M} is O, CO, SO₂, NR_J, (CO)NR_H, NR_H (CO), NR_H (SO₂), (SO₂)NR_H, or $\underline{CH_2}$;

E is $(CH_2)_{1-4}$ or $(CH_2)_m$ $O(CH_2)_p$ where $1 \le (each of m and p) \le 3$ and $2 \le (m+p) \le 4$; or E is absent;

G is R_K , OR_I or NR_JR_K , provided that if p = 1, then G is H;

R₇ is H, C₁₋₄ alkyl, C₂₋₄ alkenyl, C₂₋₄ alkynyl, C₃₋₆ cycloalkyl, phenyl, 2-pyridyl, 3-pyridyl, 4-pyridyl, (CH₂)₁₋₂Ar, where Ar is phenyl, 2-pyridyl, 3-pyridyl, or 4-pyridyl,

 $SO_2NR_H(CH_2)_{2-4}NR_JR_K$, $(CO)(CH_2)_{2-4}NR_JR_K$ or $(CO)NR_H(CH_2)_{2-4}NR_JR_K$;

 X_1 is O, S, NR_8 , or CHR_9 ; X_2 is O, S, or CHR_9 ; and X_3 is O or S; where if X_1 or X_2 is CHR_9 , said compound may also be a tautomerized indole;

 R_8 is H, C ₁₋₄ alkyl, phenyl, 2-pyridyl, 3-pyridyl, 4-pyridyl, $(CH_2)_{1-2}Ar$, where Ar is phenyl, 2-pyridyl, 3-pyridyl, or 4-pyridyl, C ₂₋₄ alkenyl, C ₂₋₄ alkynyl, C ₃₋₆ cycloalkyl, or $(C_{2-4}$ alkyl)NR_LR_M; provided R₇ and R₈ together have no more than 14 carbon atoms, exclusive of R_L, R_M, R_J and R_K;

R_G is C₁₋₄ alkyl, phenyl, 2-pyridyl, 3-pyridyl, 4-pyridyl, C₃₋₄ alkenyl, C₃₋₄ alkynyl, C₃₋₆ cycloalkyl, (CO)OR_P, (C₂₋₄ alkyl)NR_LR_M, (CO)NR_N(CH₂)₂.

4NR_LR_M, (CO)NR_LR_M, (CO)(CH₂)₂₋₄-NR_LR_M, or (CH₂)₁₋₂Ar, where Ar is phenyl, 2-pyridyl, 3-pyridyl, or 4-pyridyl;

R₉ is C₁₋₄ alkyl, phenyl, 2-pyridyl, 3-pyridyl, 4-pyridyl, C₂₋₄ alkenyl, C₂₋₄ alkynyl,

C 3.6 cycloalkyl, (CO)OR_P, (C 2.4 alkyl)NR_LR_M, (CO)NR_N(CH₂)_{2.4}NR_LR_M, (CO)NR_LR_M, (CO)(CH₂)_{2.4}-NR_LR_M, or (CH₂)_{1.2}Ar', where Ar' is phenyl, 2-pyridyl, 3-pyridyl, or 4-pyridyl;

R_P is H, C₁₋₆ alkyl, phenyl, C₃₋₄ alkenyl, C₃₋₄ alkynyl, C₃₋₆ cycloalkyl, or (CH₂)₂₋₄ NR₁R_M;

R₁₀ is H, methyl, halo, or NO₂;

R₁₁ is H, methyl, halo, or NO₂;

each of R_C, R_D, R_E, R_F, R_I, R_J, R_K, R_L and R_M is independently selected from H, C₁₋₄ alkyl, C₃₋₄ alkenyl, C₃₋₄ alkynyl, C₃₋₆ cycloalkyl, and phenyl; each of NR_CR_D,NR_ER_F, NR_JR_K, and NR_LR_M can also independently be morpholinyl, piperazinyl, pyrrolidinyl, or piperidinyl; and

each of R_H, R_N, and R_O is independently H, methyl, or ethyl;

wherein each hydrocarbon radical or heterocyclic radical above is optionally substituted with between 1 and 3 substituents independently selected from halo, C₁₋₄ alkyl, C₃₋₆ cycloalkyl, C₃₋₄ alkenyl, C₃₋₄ alkynyl, phenyl, hydroxyl, amino, (amino)sulfonyl, and NO₂, wherein each substituent alkyl, cycloalkyl, alkenyl, alkynyl or phenyl is in turn optionally substituted with between 1 and 3 substituents independently selected from halo, C₁₋₂ alkyl, hydroxyl, amino, and NO₂;

or a pharmaceutically acceptable salt or C₁₋₇ ester thereof.

4 (currently amended). The method of claim 3, wherein said neuropathic pain is associated with one of the following: inflammation, postoperative pain, phantom limb pain, burn pain, gout, trigeminal neuralgia, acute herpetic and postherpetic pain, causalgia, diabetic neuropathy, plexus avulsion, neuroma, vasculitis, viral infection, crush injury, constriction injury, tissue injury, limb amputation, post-operative pain, and arthritis pain, and any other nerve injury between the peripheral nervous system and the central nervous system, inclusively.

5 and 6 (canceled).

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7 (currently amended). The method of claim 1 A method for treating chronic pain, wherein said chronic pain is associated with inflammation, said method comprising administering to a subject in need of such treatment a composition comprising a MEK inhibitor selected from a compound of the following formula (I):

$$\mathbf{W} \stackrel{\mathbf{O}}{\longrightarrow} \mathbf{R}_{11} \stackrel{\mathbf{R}_{10}}{\longrightarrow} \mathbf{I}$$

wherein-

W is OR_1 , NR_2OR_1 , NR_AR_B , $NR_2NR_AR_B$, $O(CH_2)_{2-4}NR_AR_B$, or $NR_2(CH_2)_{2-4}NR_AR_B$;

R₁ is H, C₁₋₈ alkyl, C₃₋₈ alkenyl, C₃₋₈ alkynyl, C₃₋₈ cycloalkyl, phenyl, (phenyl)-C₁₋₄ alkyl, (phenyl)C₃₋₄ alkenyl, (phenyl)C₃₋₄ alkynyl, (C₃₋₈ cycloalkyl)C₁₋₄ alkyl, (C₃₋₈ cycloalkyl)C₃₋₄ alkenyl, (C₃₋₈ cycloalkyl)C₃₋₄ alkynyl, C₃₋₈ heterocyclic radical)C₁₋₄ alkyl, (C₃₋₈ heterocyclic radical)C₃₋₈ heterocyclic radical)C₃₋₄ alkynyl or (CH₂)₂₋₄ NR_CR_D;

R₂ is H, C₁₋₄ alkyl, phenyl, C₃₋₆ cycloalkyl, C₃₋₆ heterocyclic radical, or (C₃₋₆ cycloalkyl) methyl;

R_A is H, C₁₋₆ alkyl, C₃₋₈ alkenyl, C₃₋₈ alkynyl, C₃₋₈ cycloalkyl, phenyl, (C₃₋₈ cycloalkyl)C₁₋₄ alkyl, (C₃₋₈ cycloalkyl)C₃₋₄ alkenyl, (C₃₋₈ cycloalkyl)C₃₋₄ alkynyl, C₃₋₈ heterocyclic radical, (C₃₋₈ heterocyclic radical)C₁₋₄ alkyl,

(aminosulfonyl)phenyl, [(aminosulfonyl)phenyl]C₁₋₄ alkyl, (aminosulfonyl)C₁₋₆ alkyl, (aminosulfonyl)C₃₋₆ cycloalkyl, [(aminosulfonyl)C₃₋₆ cycloalkyl]C₁₋₄ alkyl, or (CH₂)₂₋₄ NR_CR_D;

R_B is H, C₁₋₈ alkyl, C₃₋₈ alkenyl, C₃₋₈ alkynyl, C₃₋₈ cycloalkyl, or phenyl;

Q is one of the following formulae (i) – (iii):

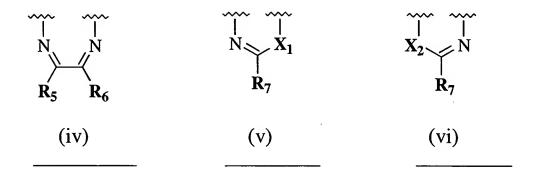
$$R_3$$
 R_4
 R_4

R_3 is H or F;

R₄ is halo, NO₂, SO₂NR_O(CH₂)₂₋₄NR_ER_F, SO₂NR_ER_F, or (CO)T;

<u>T is C₁₋₈ alkyl, C₃₋₈ cycloalkyl, (NR_ER_F)C₁₋₄ alkyl, OR_F, -NR_O(CH₂)₂₋₄ NR_ER_F, or NR_ER_F;</u>

Z is one of the following formulae (iv) – (viii):



one of R_5 and R_6 is H or methyl and the other of R_5 and R_6 is H, C_{1-6} alkyl, C_{2-6} alkenyl, C_{2-6} alkynyl, phenyl, benzyl, or -M-E-G;

 \underline{M} is O, CO, SO₂, NR_J, (CO)NR_H, NR_H (CO), NR_H (SO₂), (SO₂)NR_H, or CH₂;

<u>E is $(CH_2)_{1-4}$ or $(CH_2)_m$ O $(CH_2)_p$ where $1 \le (each of m and p) \le 3$ and $2 \le (m+p) \le 4$; or E is absent;</u>

 \underline{G} is $\underline{R}_{\underline{K}}$, $\underline{OR}_{\underline{I}}$ or $\underline{NR}_{\underline{I}}\underline{R}_{\underline{K}}$, provided that if $\underline{p} = 1$, then \underline{G} is \underline{H} ;

R₇ is H, C₁₋₄ alkyl, C₂₋₄ alkenyl, C₂₋₄ alkynyl, C₃₋₆ cycloalkyl, phenyl, 2-pyridyl, 3-pyridyl, 4-pyridyl, (CH₂)₁₋₂Ar, where Ar is phenyl, 2-pyridyl, 3-pyridyl, or 4-pyridyl,

 $SO_2NR_H(CH_2)_{2-4}NR_JR_K$, $(CO)(CH_2)_{2-4}NR_JR_K$ or $(CO)NR_H(CH_2)_{2-4}NR_JR_K$;

 X_1 is O, S, NR_8 , or CHR_9 ; X_2 is O, S, or CHR_9 ; and X_3 is O or S; where if X_1 or X_2 is CHR_9 , said compound may also be a tautomerized indole;

R₈ is H, C₁₋₄ alkyl, phenyl, 2-pyridyl, 3-pyridyl, 4-pyridyl, $(CH_2)_{1-2}Ar$, where Ar is phenyl, 2-pyridyl, or 4-pyridyl, C₂₋₄ alkenyl, C₂₋₄ alkynyl, C₃₋₆ cycloalkyl, or $(C_{2-4} \text{ alkyl})NR_LR_M$; provided R₇ and R₈ together have no more than 14 carbon atoms, exclusive of R_L, R_M, R_J and R_K;

R_G is C₁₋₄ alkyl, phenyl, 2-pyridyl, 3-pyridyl, 4-pyridyl, C₃₋₄ alkenyl, C₃₋₄ alkenyl, C₃₋₆ cycloalkyl, (CO)OR_P, (C₂₋₄ alkyl)NR_LR_M, (CO)NR_N(CH₂)₂₋₄ 4NR_LR_M, (CO)NR_LR_M, (CO)(CH₂)₂₋₄ -NR_LR_M, or (CH₂)₁₋₂Ar, where Ar is phenyl, 2-pyridyl, 3-pyridyl, or 4-pyridyl;

R₉ is C₁₋₄ alkyl, phenyl, 2-pyridyl, 3-pyridyl, 4-pyridyl, C₂₋₄ alkenyl, C₂₋₄ alkynyl,

C 3.6 cycloalkyl, (CO)OR_P, (C 2.4 alkyl)NR₁R_M, (CO)NR_N(CH₂)₂₋₄NR₁R_M, (CO)NR_LR_M, (CO)(CH₂)₂₋₄-NR_LR_M, or (CH₂)₁₋₂Ar', where Ar' is phenyl, 2-pyridyl, 3-pyridyl, or 4-pyridyl;

R_P is H, C₁₋₆ alkyl, phenyl, C₃₋₄ alkenyl, C₃₋₄ alkynyl, C₃₋₆ cycloalkyl, or (CH₂)₂₋₄ NR_LR_M;

R₁₀ is H, methyl, halo, or NO₂;

R₁₁ is H, methyl, halo, or NO₂;

each of R_C , R_D , R_E , R_F , R_I , R_I , R_K , R_L and R_M is independently selected from H, C_{1-4} alkyl, C_{3-4} alkenyl, C_{3-4} alkynyl, C_{3-6} cycloalkyl, and phenyl; each of NR_CR_D , NR_ER_F , NR_IR_K , and NR_LR_M can also independently be morpholinyl, piperazinyl, pyrrolidinyl, or piperidinyl; and

each of R_H, R_N, and R_O is independently H, methyl, or ethyl;

wherein each hydrocarbon radical or heterocyclic radical above is optionally substituted with between 1 and 3 substituents independently selected from halo, C₁₋₄ alkyl, C₃₋₆ cycloalkyl, C₃₋₄ alkenyl, C₃₋₄ alkynyl, phenyl, hydroxyl, amino, (amino)sulfonyl, and NO₂, wherein each substituent alkyl, cycloalkyl, alkenyl, alkynyl or phenyl is in turn optionally substituted with between 1 and 3 substituents independently selected from halo, C₁₋₂ alkyl, hydroxyl, amino, and NO₂;

or a pharmaceutically acceptable salt or C₁₋₇ ester thereof.

8 (currently amended). The method of claim 1 A method for treating chronic pain, wherein said chronic pain is associated with arthritis, said method comprising administering to a subject in need of such treatment a composition comprising a MEK inhibitor selected from a compound of the following formula (I):

wherein

W is OR_1 , NR_2OR_1 , NR_AR_B , $NR_2NR_AR_B$, $O(CH_2)_{2-4}NR_AR_B$, or $NR_2(CH_2)_{2-4}NR_AR_B$;

R₁ is H, C₁₋₈ alkyl, C₃₋₈ alkenyl, C₃₋₈ alkynyl, C₃₋₈ cycloalkyl, phenyl, (phenyl)-C₁₋₄ alkyl, (phenyl)C₃₋₄ alkenyl, (phenyl)C₃₋₄ alkynyl, (C₃₋₈ cycloalkyl)C₁₋₄ alkyl, (C₃₋₈ cycloalkyl)C₃₋₄ alkenyl, (C₃₋₈ cycloalkyl)C₃₋₄ alkynyl, C₃₋₈ heterocyclic radical)C₁₋₄ alkyl, (C₃₋₈ heterocyclic radical)C₃₋₄ alkynyl or (CH₂)₂₋₄ NR_CR_D;

R₂ is H, C₁₋₄ alkyl, phenyl, C₃₋₆ cycloalkyl, C₃₋₆ heterocyclic radical, or (C₃₋₆ cycloalkyl) methyl;

R_A is H, C₁₋₆ alkyl, C₃₋₈ alkenyl, C₃₋₈ alkynyl, C₃₋₈ cycloalkyl, phenyl, (C₃₋₈ cycloalkyl)C₁₋₄ alkyl, (C₃₋₈ cycloalkyl)C₃₋₄ alkenyl, (C₃₋₈ cycloalkyl)C₃₋₄ alkynyl, C₃₋₈ heterocyclic radical, (C₃₋₈ heterocyclic radical)C₁₋₄ alkyl, (aminosulfonyl)phenyl, [(aminosulfonyl)phenyl]C₁₋₄ alkyl, (aminosulfonyl)C₁₋₆ alkyl, (aminosulfonyl)C₃₋₆ cycloalkyl, [(aminosulfonyl)C₃₋₆ cycloalkyl]C₁₋₄ alkyl, or (CH₂)₂₋₄ NR_CR_D;

R_B is H, C₁₋₈ alkyl, C₃₋₈ alkenyl, C₃₋₈ alkynyl, C₃₋₈ cycloalkyl, or phenyl;

Q is one of the following formulae (i) – (iii):

R₃ is H or F;

R4 is halo, NO₂, SO₂NR_O(CH₂)₂₋₄NR_ER_F, SO₂NR_ER_F, or (CO)T;

<u>T is C₁₋₈ alkyl, C₃₋₈ cycloalkyl, (NR_ER_F)C₁₋₄ alkyl, OR_F, -NR_O(CH₂)₂₋₄ NR_ER_F, or NR_ER_F;</u>

Z is one of the following formulae (iv) – (viii):

one of R_5 and R_6 is H or methyl and the other of R_5 and R_6 is H, C_{1-6} alkyl, C_{2-6} alkenyl, C_{2-6} alkynyl, phenyl, benzyl, or -M-E-G;

 \underline{M} is O, CO, SO₂, NR_J, (CO)NR_H, NR_H (CO), NR_H (SO₂), (SO₂)NR_H, or CH₂;

E is $(CH_2)_{1-4}$ or $(CH_2)_m$ $O(CH_2)_p$ where $1 \le (\text{each of m and p}) \le 3$ and $2 \le (m+p) \le 4$; or E is absent;

G is R_K , OR_I or NR_JR_K , provided that if p = 1, then G is H;

R₇ is H, C₁₋₄ alkyl, C₂₋₄ alkenyl, C₂₋₄ alkynyl, C₃₋₆ cycloalkyl, phenyl, 2-pyridyl, 3-pyridyl, 4-pyridyl, (CH₂)₁₋₂Ar, where Ar is phenyl, 2-pyridyl, 3-pyridyl, or 4-pyridyl,

 $SO_2NR_H(CH_2)_2$ $4NR_JR_K$, $(CO)(CH_2)_2$ $4NR_JR_K$ or $(CO)NR_H(CH_2)_2$ $4NR_JR_K$;

 X_1 is O, S, NR_8 , or CHR_9 ; X_2 is O, S, or CHR_9 ; and X_3 is O or S; where if X_1 or X_2 is CHR_9 , said compound may also be a tautomerized indole;

 R_8 is H, C₁₋₄ alkyl, phenyl, 2-pyridyl, 3-pyridyl, 4-pyridyl, $(CH_2)_{1-2}Ar$, where Ar is phenyl, 2-pyridyl, 3-pyridyl, or 4-pyridyl, C₂₋₄ alkenyl, C₂₋₄ alkynyl, C₃₋₆ cycloalkyl, or $(C_{2-4}$ alkyl)NR_LR_M; provided R₇ and R₈ together have no more than 14 carbon atoms, exclusive of R_L, R_M, R_J and R_K;

R_G is C₁₋₄ alkyl, phenyl, 2-pyridyl, 3-pyridyl, 4-pyridyl, C₃₋₄ alkenyl, C₃₋₄ alkynyl, C₃₋₆ cycloalkyl, (CO)OR_P, (C₂₋₄ alkyl)NR_LR_M, (CO)NR_N(CH₂)₂₋₄ 4NR_LR_M, (CO)NR_LR_M, (CO)(CH₂)₂₋₄ -NR_LR_M, or (CH₂)₁₋₂Ar, where Ar is phenyl, 2-pyridyl, 3-pyridyl, or 4-pyridyl;

R₉ is C₁₋₄ alkyl, phenyl, 2-pyridyl, 3-pyridyl, 4-pyridyl, C₂₋₄ alkenyl, C₂₋₄ alkynyl,

C₃₋₆ cycloalkyl, (CO)OR_P, (C₂₋₄ alkyl)NR_LR_M, (CO)NR_N(CH₂)₂₋₄NR_LR_M, (CO)NR_LR_M, (CO)(CH₂)₂₋₄-NR_LR_M, or (CH₂)₁₋₂Ar', where Ar' is phenyl, 2-pyridyl, 3-pyridyl, or 4-pyridyl;

 R_P is H, C₁₋₆ alkyl, phenyl, C₃₋₄ alkenyl, C₃₋₄ alkynyl, C₃₋₆ cycloalkyl, or $(CH_2)_{2-4}$ NR_LR_M ;

R₁₀ is H, methyl, halo, or NO₂;

R₁₁ is H, methyl, halo, or NO₂;

each of R_C, R_D, R_E, R_F, R_I, R_J, R_K, R_L and R_M is independently selected from H, C₁₋₄ alkyl, C₃₋₄ alkenyl, C₃₋₄ alkynyl, C₃₋₆ cycloalkyl, and phenyl; each of NR_CR_D,NR_ER_F, NR_JR_K, and NR_LR_M can also independently be morpholinyl, piperazinyl, pyrrolidinyl, or piperidinyl; and

each of R_H, R_N, and R_O is independently H, methyl, or ethyl;

wherein each hydrocarbon radical or heterocyclic radical above is optionally substituted with between 1 and 3 substituents independently selected from halo, C₁₋₄ alkyl, C₃₋₆ cycloalkyl, C₃₋₄ alkenyl, C₃₋₄ alkynyl, phenyl, hydroxyl, amino, (amino)sulfonyl, and NO₂, wherein each substituent alkyl, cycloalkyl, alkenyl, alkynyl or phenyl is in turn optionally substituted with between 1 and 3 substituents independently selected from halo, C₁₋₂ alkyl, hydroxyl, amino, and NO₂;

or a pharmaceutically acceptable salt or C₁₋₇ ester thereof.

9 (currently amended). The method of claim 1 A method for treating chronic pain, wherein said chronic pain is associated with post-operative pain, said method comprising administering to a subject in need of such treatment a composition comprising a MEK inhibitor selected from a compound of the following formula (I):

$$\begin{array}{c}
O \\
M \\
N
\end{array}$$

$$\begin{array}{c}
H \\
R_{10} \\
R_{11}
\end{array}$$

$$I$$

wherein

W is OR_1 , NR_2OR_1 , NR_AR_B , $NR_2NR_AR_B$, $O(CH_2)_{2-4}NR_AR_B$, or $NR_2(CH_2)_{2-4}$ NR_AR_B ;

R₁ is H, C ₁₋₈ alkyl, C ₃₋₈ alkenyl, C ₃₋₈ alkynyl, C ₃₋₈ cycloalkyl, phenyl, (phenyl)-C ₁₋₄ alkyl, (phenyl)C ₃₋₄ alkenyl, (phenyl)C ₃₋₄ alkynyl, (C ₃₋₈ cycloalkyl)C ₁₋₄ alkyl, (C ₃₋₈ cycloalkyl)C ₃₋₄ alkenyl, (C ₃₋₈ cycloalkyl)C ₃₋₄ alkynyl, C ₃₋₈ heterocyclic radical)C ₁₋₄ alkyl, (C ₃₋₈ heterocyclic radical)C ₁₋₄ alkyl, (C ₃₋₈ heterocyclic radical)C ₃₋₄ alkynyl or (CH₂)₂₋₄ NR_CR_D;

 R_2 is H, C _{1.4} alkyl, phenyl, C _{3.6} cycloalkyl, C _{3.6} heterocyclic radical, or (C _{3.6} cycloalkyl) methyl;

R_A is H, C₁₋₆ alkyl, C₃₋₈ alkenyl, C₃₋₈ alkynyl, C₃₋₈ cycloalkyl, phenyl, (C₃₋₈ cycloalkyl)C₁₋₄ alkyl, (C₃₋₈ cycloalkyl)C₃₋₄ alkenyl, (C₃₋₈ cycloalkyl)C₃₋₄ alkynyl, C₃₋₈ heterocyclic radical, (C₃₋₈ heterocyclic radical)C₁₋₄ alkyl, (aminosulfonyl)phenyl, [(aminosulfonyl)phenyl]C₁₋₄ alkyl, (aminosulfonyl)C₁₋₆ alkyl, (aminosulfonyl)C₃₋₆ cycloalkyl, [(aminosulfonyl)C₃₋₆ cycloalkyl]C₁₋₄ alkyl, or (CH₂)₂₋₄ NR_CR_D;

R_B is H, C₁₋₈ alkyl, C₃₋₈ alkenyl, C₃₋₈ alkynyl, C₃₋₈ cycloalkyl, or phenyl;

Q is one of the following formulae (i) – (iii):

$$R_3$$
 R_4
 R_4

R_3 is H or F;

R4 is halo, NO2, SO2NRO(CH2)2.4NRERF, SO2NRERF, or (CO)T;

 $\frac{\text{T is C}_{1-8} \text{ alkyl, C}_{3-8} \text{ cycloalkyl, (NR}_{E}R_{F})C_{1-4} \text{ alkyl, OR}_{F}, -\text{NR}_{O}(\text{CH}_{2})_{2-4} \text{ NR}_{E}R_{F},}{\text{or NR}_{E}R_{F};}$

Z is one of the following formulae (iv) – (viii):

$$\begin{array}{ccc}
& & & & & \\
& & & & \\
N & & & & \\
N & & & \\

\end{array}$$
(viii) (viii)

one of R₅ and R₆ is H or methyl and the other of R₅ and R₆ is H, C₁₋₆ alkyl, C₂₋₆ alkenyl, C₂₋₆ alkynyl, phenyl, benzyl, or -M-E-G;

 \underline{M} is O, CO, SO₂, NR_J, (CO)NR_H, NR_H (CO), NR_H (SO₂), (SO₂)NR_H, or CH₂;

<u>E is $(CH_2)_{1-4}$ or $(CH_2)_m$ $O(CH_2)_p$ where $1 \le (each of m and p) \le 3$ and $2 \le (m+p) \le 4$; or E is absent;</u>

G is R_K , OR_I or NR_JR_K , provided that if p = 1, then G is H;

R₇ is H, C₁₋₄ alkyl, C₂₋₄ alkenyl, C₂₋₄ alkynyl, C₃₋₆ cycloalkyl, phenyl, 2-pyridyl, 3-pyridyl, 4-pyridyl, (CH₂)₁₋₂Ar, where Ar is phenyl, 2-pyridyl, 3-pyridyl, or 4-pyridyl,

 $SO_2NR_H(CH_2)_{2-4}NR_JR_K$, $(CO)(CH_2)_{2-4}NR_JR_K$ or $(CO)NR_H(CH_2)_{2-4}NR_JR_K$;

 X_1 is O, S, NR_8 , or CHR_9 ; X_2 is O, S, or CHR_9 ; and X_3 is O or S; where if X_1 or X_2 is CHR_9 , said compound may also be a tautomerized indole;

R₈ is H, C₁₋₄ alkyl, phenyl, 2-pyridyl, 3-pyridyl, 4-pyridyl, $(CH_2)_{1-2}Ar$, where Ar is phenyl, 2-pyridyl, 3-pyridyl, or 4-pyridyl, C₂₋₄ alkenyl, C₂₋₄ alkynyl, C₃₋₆ cycloalkyl, or $(C_{2-4} \text{ alkyl})NR_LR_M$; provided R₇ and R₈ together have no more than 14 carbon atoms, exclusive of R_L, R_M, R_J and R_K;

R_G is C₁₋₄ alkyl, phenyl, 2-pyridyl, 3-pyridyl, 4-pyridyl, C₃₋₄ alkenyl, C₃₋₄ alkynyl, C₃₋₆ cycloalkyl, (CO)OR_P, (C₂₋₄ alkyl)NR_LR_M, (CO)NR_N(CH₂)₂₋
4NR_LR_M, (CO)NR_LR_M, (CO)(CH₂)₂₋₄-NR_LR_M, or (CH₂)₁₋₂Ar, where Ar is phenyl, 2-pyridyl, 3-pyridyl, or 4-pyridyl;

R₉ is C₁₋₄ alkyl, phenyl, 2-pyridyl, 3-pyridyl, 4-pyridyl, C₂₋₄ alkenyl, C₂₋₄ alkynyl,

C 3.6 cycloalkyl, (CO)OR_P, (C 2.4 alkyl)NR_LR_M, (CO)NR_N(CH₂)_{2.4}NR_LR_M, (CO)NR_LR_M, (CO)(CH₂)_{2.4}-NR_LR_M, or (CH₂)_{1.2}Ar', where Ar' is phenyl, 2-pyridyl, 3-pyridyl, or 4-pyridyl;

R_P is H, C₁₋₆ alkyl, phenyl, C₃₋₄ alkenyl, C₃₋₄ alkynyl, C₃₋₆ cycloalkyl, or (CH₂)₂₋₄ NR_LR_M;

R₁₀ is H, methyl, halo, or NO₂;

R₁₁ is H, methyl, halo, or NO₂;

each of R_C, R_D, R_E, R_F, R_I, R_J, R_K, R_L and R_M is independently selected from H, C₁₋₄ alkyl, C₃₋₄ alkenyl, C₃₋₄ alkynyl, C₃₋₆ cycloalkyl, and phenyl; each of NR_CR_D,NR_ER_F, NR_JR_K, and NR_LR_M can also independently be morpholinyl, piperazinyl, pyrrolidinyl, or piperidinyl; and

each of R_H, R_N, and R_O is independently H, methyl, or ethyl;

wherein each hydrocarbon radical or heterocyclic radical above is optionally substituted with between 1 and 3 substituents independently selected from halo, C₁₋₄ alkyl, C₃₋₆ cycloalkyl, C₃₋₄ alkenyl, C₃₋₄ alkynyl, phenyl, hydroxyl, amino, (amino)sulfonyl, and NO₂, wherein each substituent alkyl, cycloalkyl, alkenyl,

alkynyl or phenyl is in turn optionally substituted with between 1 and 3 substituents independently selected from halo, C₁₋₂ alkyl, hydroxyl, amino, and NO₂;

or a pharmaceutically acceptable salt or C_{1.7} ester thereof.

10 (currently amended). A method of claim 1 claim 8, wherein Q is formula (i).

11 (original). A method of claim 10, wherein R_3 is H or fluoro.

12 (original). A method of claim 11, wherein R₄ is fluoro, chloro, or bromo.

13 (currently amended). A method of elaim 1 claim 8, wherein R_{10} is hydrogen, methyl, fluoro, or chloro.

14 (currently amended). A method of claim 1 claim 8, wherein R_{11} is methyl, chloro, fluoro, nitro, or hydrogen.

15 (original). A method of claim 14, wherein R_{11} is H.

16 (original). A method of claim 14, wherein R_{11} is fluoro.

17 (original). A method of claim 13, wherein each of R_{10} and R_{11} is fluoro.

18 (currently amended). A method of claim 1 claim 8, wherein R₁ is H, methyl, ethyl, propyl, isopropyl, isobutyl, benzyl, phenethyl, allyl, C ₃₋₅ alkenyl, C ₃₋₆ cycloalkyl, (C ₃₋₅ cycloalkyl)C ₁₋₂ alkyl, (C ₃₋₅ heterocyclic radical)C ₁₋₂ alkyl, or (CH₂)₂₋₄ NR_CR_D.

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19 (original). A method of claim 18, wherein R_1 is H or (C $_{3-4}$ cycloalkyl)C $_{1-2}$ alkyl.

20 (currently amended). A method of claim 1 claim 8, wherein R_2 is H or methyl.

21 (currently amended). A method of claim 1 claim 8, wherein R_A has at least one hydroxyl substituent.

22 (currently amended). A compound of elaim 1 claim 8, wherein R_A is H, methyl, ethyl, isobutyl, hydroxyethyl, phenyl, 2-piperidin-1-yl-ethyl, 2,3-dihydroxy-propyl, 3-[4-(2-hydroxyethyl)-piperazin-1-yl]-propyl, 2-pyrrolidin-1-yl-ethyl, or 2-diethylamino-ethyl; and R_B is H; or where R_B is methyl and R_A is phenyl.

23 (currently amended). A method of elaim 1 claim 8, wherein W is NR_AR_B or $NR_2NR_AR_B$.

24 (currently amended). A method of elaim-1 claim 8, wherein W is $NR_2(CH_2)_{2-4} NR_A R_B$ or $O(CH_2)_{2-3} NR_A R_B$.

25 (currently amended). A method of elaim 1 claim 8, wherein W is NR_2OR_1 .

26 (currently amended). A method of elaim 1 claim 8, wherein W is OR_1 .

27 (currently amended). A method of claim 1 claim 8, wherein Z is formula (v).

28 (original). A method of claim 27, wherein X₁ is NR₈, and R₇ is H.

29 (currently amended). A method of elaim 1 claim 8, wherein said MEK inhibitor has a structure selected from: 7-fluoro-6-(4-iodo-2-methyl-phenylamino)-1H-benzoimidazole-5-carboxylic acid.

30 (currently amended). A method of elaim 1 claim 8, wherein said MEK inhibitor has a structure selected from: 7-fluoro-6-(4-iodo-2-methyl-phenylamino)-1H-benzoimidazole-5-carboxylic acid; 7-fluoro-6-(4-iodo-2-methyl-phenylamino)-benzooxazole-5-carboxylic acid; 7-fluoro-6-(4-iodo-2-methyl-phenylamino)-benzothiazole-5-carboxylic acid; 7-fluoro-6-(4-iodo-2-methyl-phenylamino)-benzo[1,2,5]thiadiazole-5-carboxylic acid; 7-fluoro-6-(4-iodo-2-methyl-phenylamino)-benzo[1,2,5]oxadiazole-5-carboxylic acid; 7-fluoro-6-(4-iodo-2-methyl-phenylamino)-2-(2-hydroxyethyl)-1H-benzoimidazole-5-carboxylic acid; 7-fluoro-6-(4-iodo-2-methyl)-1H-benzoimidazole-5-carboxylic acid; 7-fluoro-6-(4-iodo-2-methyl)-1H-benzoimidazole-5-carboxylic acid; 8-fluoro-7-(4-iodo-2-methyl-phenylamino)-quinoxaline-6-carboxylic acid; and 7-fluoro-6-(4-iodo-2-methyl-phenylamino)-1H-benzotriazole-5-carboxylic acid; and the corresponding hydroxamic acids and cyclopropylmethyl hydroxamates.

31 (currently amended). The method of elaim 1 claim 8 wherein said MEK inhibitor has a structure selected from: 7-fluoro-6-(4-iodo-2-methyl-phenylamino)-1*H*-benzoimidazole-5-carboxylic acid cyclopropylmethoxy-amide; 7-fluoro-6-(4-iodo-2-methyl-phenylamino)-6,7-dihydro-1*H*-benzoimidazole-5-carboxylic acid (hydrochloride); 7-fluoro-6-(4-iodo-2-methyl-phenylamino)-1*H*-benzoimidazole-5-carboxylic acid; 7-fluoro-6-(4-iodo-2-methyl-phenylamino)-3*H*-benzoimidazole-5-carboxylic acid (2-hydroxy-ethoxy)-amide; 6-(2-chloro-4-iodo-phenylamino)-7-fluoro-1*H*-benzoimidazole-5-carboxylic acid; and 7-fluoro-6-(4-iodo-2-methyl-phenylamino)-1*H*-benzoimidazole-5-carboxylic acid pentafluorophenyl ester.

32 (currently amended). The method of elaim-1 claim 8 wherein said MEK inhibitor has a structure selected from: 7-fluoro-6-(4-iodo-2-methyl-phenylamino)-1*H*-benzoimidazole-5-carboxylic acid cyclopropylmethoxy-amide; and 7-fluoro-6-(4-iodo-2-methyl-phenylamino)-3*H*-benzoimidazole-5-carboxylic acid (2-hydroxy-ethoxy)-amide.